```
Post-processing: Minimum Match 0%
Maximum Match 100%
Listing first 45 summaries
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Database :
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Minimum DB seq length: 0
Maximum DB seq length: 2000000000
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Total number of hits satisfying chosen parameters:
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Scoring table:
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Sequence:
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Run on:
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    OM nucleic - nucleic search, using sw model
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Perfect score:
                                                                                                                                                                                                                                                                                                                           Result
                                                                                                                                                                                                                                                                                                        No.
                                                                                                                                                                                                                                                                                                                                                                                                                  Pred. No. is the number of results predicted by chance to have a
score greater than or equal to the score of the result being printed,
and is derived by analysis of the total score distribution.
                                                                                                            470.6
470.6
470.6
470.6
470.6
470.6
470.6
469
467.4
400.6
279.8
268.4
                                                                                                                                                                                                                                                                                                     Score
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   N_Geneseq_36:*

1: /SIDS6/gcgdata/geneseq/geneseqn/NA1981.DAT:*
2: /SIDS6/gcgdata/geneseq/geneseqn/NA1981.DAT:*
3: /SIDS6/gcgdata/geneseq/geneseqn/NA1981.DAT:*
4: /SIDS6/gcgdata/geneseq/geneseqn/NA1982.DAT:*
5: /SIDS6/gcgdata/geneseq/geneseqn/NA1983.DAT:*
5: /SIDS6/gcgdata/geneseq/geneseqn/NA1984.DAT:*
6: /SIDS6/gcgdata/geneseq/geneseqn/NA1985.DAT:*
7: /SIDS6/gcgdata/geneseq/geneseqn/NA1986.DAT:*
7: /SIDS6/gcgdata/geneseq/geneseqn/NA1989.DAT:*
9: /SIDS6/gcgdata/geneseq/geneseqn/NA1989.DAT:*
10: /SIDS6/gcgdata/geneseq/geneseqn/NA1999.DAT:*
11: /SIDS6/gcgdata/geneseq/geneseqn/NA199.DAT:*
12: /SIDS6/gcgdata/geneseq/geneseqn/NA199.DAT:*
13: /SIDS6/gcgdata/geneseq/geneseqn/NA199.DAT:*
14: /SIDS6/gcgdata/geneseq/geneseqn/NA199.DAT:*
15: /SIDS6/gcgdata/geneseq/geneseqn/NA199.DAT:*
16: /SIDS6/gcgdata/geneseq/geneseqn/NA199.DAT:*
17: /SIDS6/gcgdata/geneseq/geneseqn/NA199.DAT:*
18: /SIDS6/gcgdata/geneseq/geneseqn/NA199.DAT:*
19: /SIDS6/gcgdata/geneseq/geneseqn/NA199.DAT:*
                                                                                                                                                                                                                                                                                                        Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       November 25, 2000, 04:36:10 ; Search time 88 Seconds (without alignments) 2010.649 Million cell updates/sec
480022 seqs, 187831343 residues
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Gapop 10.0 , Gapext 1.0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                1 AACTITGGCCGACTTCACTG.....TCACTAACTTACATCAAAGT 471
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            US-09-373-230-1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        GenCore version 4.5
Copyright (c) 1993 - 2000 Comp
                                                                                                                                                                                                                                                                                                        Length DB
                                                                   471
471
471
471
471
471
471
471
471
                                                                                                                                                                                                                                                                                                        ij
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               1.,
                                                                                                                                                                                                                                                         T32403
                                                                                                                                                                                                                                                                                                                                                                            SUMMARIES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Compugen Ltd
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                960044
                DNA encoding a pro
Wild-type mouse in ter
Mutant mouse inter
Mutant mouse inter
Nucleotide sequenc
Nucleotide sequenc
                                                                                                                                                                Mouse interferon-g
Interferon gamma p
Mouse interferon-g
Murine protein for
Mouse interleukin
                                                                                                                                                                                                                                                                                                        Description
Equine interleukin
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   T32403
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         RESULT
```

•	-	-			_	_	_			-																						
6.1	127.8	243	243	244.4		244.4	•		•		•	244.6			•		•	•	•	•	•	•	•	•		•	•	•		•	247.8	•
27.1	7.	51.6	۳.	۳	۳.	51.9		μ.	1		۲.	Ϊ.	1.	52.3	ν.	ν.	2	.2	٥.				.2	2						.2	52.6	
28994	11464	471	471	1427	1427	582	582	582	540	540	471	471	471	471	471	1120	1120	1120	579	579	471	471	471	471	1380	1101	1101	570	471	471	471	477
19	19	19	19										19	19	19	21	19	17	19	19	21	18	17	17	19	18	18	19	19	19	19	21
82	V48228	63	V48230	A13795	X27726	Z55623	A13793	X27724	A13801	X27732	V32630	V32629	V48229	V32628	V32627	Z36876	V15825	T32404	V17200	V18906	Z36875	T80209	T32402	T32411	V05368	æ	T74988	$\sigma$	$\sim$	62	22	A10526
Genomic DNA for in	Interleukin 18 con	Mutant human inter	Human interleukin	Canine interleukin	Recombinant canine	Mutant human inter	Mutant human inter	Human interleukin	Mutant human inter	ř	DNA encoding a pro	erfer	Human interferon-g	Interferon-gamma i	Interferon-gamma i	DNA encoding a mat	Human protein for	interferon-	Human interferon-g	cDNA encoding huma	Interferon gamma i	Interferon gamma i	Wild-type human in	Mutant human inter	Mutant human inter	_	Human interleukin-					

## ALIGNMENTS

T32403;

Mouse interferon-gamma inducer protein cDNA

29-SEP-1996 (first entry)

T32403 standard; cDNA to mRNA; 471

```
29-SEP-1995;
15-NOV-1994;
23-FEB-1995;
10-MAR-1995;
18-SEP-1995;
WPI; 1996-252837/26.
P-PSDB; R99559.
                         Fukuda S, Kohno K, Kunikata T, Kurimoto M, Taniguchi M, Tanimoto T, Torigoe K, Ushio S;
                                                                                                                                               22-MAY-1996
                                                                                                                                                                 EP712931-A2
                                                                                                                                                                                                            Interferon-gamma inducer protein; IFN-gamma; antiviral; virucide;
antitumour; antibacterial; immunoregulatory; adoptive immunotherapy;
                                                     (HAYB ) HAYASHIBARA SEIBUTSU KAGAKU.
                                                                                                                             10-NOV-1995;
                                                                                                                                                                                   Mus sp.
                                                                                                                                                                                                    therapy; cancer; ds.
                                                                               95JP-0274988.
94JP-0304203.
95JP-0058240.
95JP-0078357.
                                                                        95JP-0262062.
                                                                                                                              95EP-0308055
                                      Okamura
```

```
RESULT
T16224
ID T1
XX T1
XX T1
XX T1
XX T1
XX IN
DT 02
DT 02
XX IN
XX IN
XX IN
XX IN
XX IN
XX IN
XX AN
XX AN
                                                                                                                                                                                                                                                                                                                                                                                                               밁
                                                                                                                                                                                                                                                                                                                                                                        δÃ
                                                                                                                                                                                                                                                                                                                                                                                                                                           Qy
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  В
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   QΥ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Ωy
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          밁
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         δÃ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             밁
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            γQ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   밁
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      В
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Qy
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        В
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Query Match
Best Local
                             antitumour; antiseptic; immunoregulatory; platelet-increasing agent; therapy; prevention; condyloma acuminatum; renal cancer; brain cance granuloma; mycosis fungoides; rheumatism; allergy; cytotoxicity; AID killer T-cell; interleukin-2; IL-2; tumour necrosis factor; TNF;
              adoptive immunotherapy; monoclonal antibody; ds
                                                                                                     Interferon gamma; inducer; IFNgamma; immunocompetent cell; antiviral;
                                                                                                                                        Interferon gamma production inducer protein coding sequence.
                                                                                                                                                                             02-SEP-1996
                                                                                                                                                                                                                 T16224;
                                                                                                                                                                                                                                                  T16224 standard;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               induces interferon-gamma (IFN-gamma) prodn. by immunocompetent cells. The clone was obtd. from a mouse liver cDNA library by PCR amplification using primers (see also T32405-06) based on tryptic peptides (R99561-62) of the protein. A DNA fragment based on the cDNA clone was used to screen a human liver cDNA library, leading to the isolation of a clone (T32402) coding for human mature IFN-gamma inducer protein (R99558), a useful therapeutic agent.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Sequence 471 BP;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               A cDNA clone (T32403) codes for a novel mouse protein (R99559)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Example A-3-2;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         DNA encoding
                                                                                                                                                                                                                                                                                                                                         421
                                                                                                                                                                                                                                                                                                                                                                                                               361
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      301
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         301
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          241
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             241
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              181
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  181
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    121
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   121
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      61
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         19
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Ľ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        treat and
                                                                                                                                                                                                                                                                                     N
                                                                                                                                                                                                                                                                                                                                                                          GAAAATGGGGATAAATCTGTAATGTTCACTCTCACTAACTTACATCAAAGT 471
                                                                                                                                                                                                                                                                                                                                                                                                                            gaaaatggggataaatctgtaatgttcactctcactaacttacatcaaagt 471
                                                                                                                                                                                                                                                                                                                                                                                                            cactttcttgcttgccaaaaggaagatgatgctttcaaactcattctgaaaaaaaggat
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         TCCTTTGAGGAAATGGATCCACCTGAAAATATTGATGATATACAAAGTGATCTCATATTC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             gtgaccctctctgtgaaggatagtaaaaygtctaccctctcctgtaagaacaagatcatt
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          GTGACCCTCTGTGAAGGATAGTAAAAXGTCTACCCTCTCCTGTAAGAACAAGATCATT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  GAACCCCAGACCAGACTGATAATATACATGTACAAAGACAGTGAAGTAAGAGGACTGGCT 180
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  TTCGTTGACAAAAGACAGCCTGTGTTCGAGGATATGACTGATATTGATCAAAGTGCCAGT 120
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        AACTTTGGCCGACTTCACTGTACAACCGCAGTAATACGGGAATATAAATGACCAAGTTCTC 60
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                tttcagaaacgtgttccaggacacaacaagatggagtttgaatcttcactgtatgaagga
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   TTTCAGAAACGTGTTCCAGGACACAACAAGATGGAGTTTGAATCTTCACTGTATGAAGGA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    tcctttgaggaaatggatccacctgaaaatattgatgatatacaaagtgatctcatattc
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             gaaccccagaccagactgataatatacatgtacaaagacagtgaagtaagaggactggct
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ttcgttgacaaaagacagcctgtgttcgaggatatgactgatattgatcaaagtgccagt 120
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      aactttggccgacttcactgtacaaccgcagtaatacggaatataaatgaccaagttctc
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              471;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Conservative
                                                                                                                                                                            (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    interferon-gamma prodn.-inducing polypeptide - useful
prevent, e.g. viral disease, malignancies and immune
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Page 36-37; 48pp; English
                                                                                                                                                                                                                                                  CDNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    162 A; 91 C; 92 G; 125 T; 1 other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           100.0%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                99.98;
                                                                                                                                                                                                                                                  ç
                                                                                                                                                                                                                                                  mRNA; 471
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             0;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Score 470.6;
Pred. No. 5
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Mismatches
                                                                                                                                                                                                                                                  ВP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           5.8e-124;
                                                                  renal cancer; brain cancer;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                DB
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Length 471;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           0;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            240
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               . 180
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  360
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      60
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           0;
```

```
This sequence represents the coding sequence for the interferon gamma (IFNgamma) inducer protein of the invention. The encoded protein induc IFNgamma production in immunocompetent cells. The protein is useful as an antiviral, antitumour, antiseptic, immunoregulatory and platelet-increasing agent. It can be used for treating or preventing AIDS, condyloma acuminatum, renal or brain cancer, granuloma, mycosis fungoides, rheumatism and allergy. The protein can also be used to induce IFNgamma production in cultured cells. The IFNgamma inducer strongly induces cytotoxicity of killer Teclis and when used with interleukin-2 (IL-2) and tumour necrosis factor (TNF), may improve the effect (or reduce side effects) of adoptive immunotherapy in tumours. This sequence can be used to produce the protein, which can then be never interfied.
                                                                                                                                                                                                                                                                                        rrotein that induces gamma interferon prodn. cells - used e.g. as antiviral or antitumour cytotoxicity of killer cells
 Sequence 471 BP;
                                                                                                                                                                                                                                                                Claim 4; Page 22-23;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Mus musculus
                                                                                                                                                                                                                                                                                                                                                                   P-PSDB; R92506.
                                                                                                                                                                                                                                                                                                                                                                                                                    Tanimoto T,
                                                                                                                                                                                                                                                                                                                                                                                                                                     Kohno K,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                      (HAYB)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       14-JUL-1994;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        13-JUL-1995;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     10-FEB-1995;
                                                                                                                                                                                                                                                                                                                                                                                   1996-070177/08.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                      HAYASHIBARA SEIBUTSU KAGAKU.
                               (or assayed) using monoclonal antibodies.
                                                                                                                                                                                                                                                                                                                                                                                                                                     Kunikata T,
                                                                                                                                                                                                                                                                                                                                                                                                                  Torigoe
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      95JP-0045057.
94JP-0184162.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        95EP-0304906
162 A; 91 C;
                                                                                                                                                                                                                                                              30pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                     Kurimoto M, Okamura H,
92 G; 125
T; 1 other;
                                                                                                                                                                                                                                                                                                               in immuno:competent agent, also induces
                                                                                                                                                                                                                                                                                                                                                                                                                                  Taniguchi
                                                                                                                                                                                                               induces
                                                                                                                                                                                               as
```

Matches 471; Query Match Local Similarity Conservative 100.0%; 99.98; 0; Score 470.6; Pred. No. 5 Mismatches .8e-124; DB 17; ç Indels Length 471; 0; Gaps

0

밁 ρy 밁 δÃ 밁 Qy 밁 Qy Дb Qy 밁 δÃ. ρ B 361 301 301 181 181 121 121 61 61 TTCGTTGACAAAAGACAGCCTGTGTTCGAGGATATGACTGATATTGATCAAAGTGCCAGT 120 1 AACTTTGGCCGACTTCACTGTACAACCGCAGTAATACGGAATATAAAATGACCAAGTTCTC 1 aactttggccgacttcactgtacaaccgcagtaatacggaatataaatgaccaagttctc CACTTTCTTGCTTGCCAAAAGGAAGATGATGCTTTCAAACTCATTCTGAAAAAAAGGAT GAACCCCAGACCAGACTGATAATATACATGTACAAAGACAGTGAAGTAAGAGGACTGGCT 180 tttcagaaacgtgttccaggacacaacaagatggagtttgaatcttcactgtatgaagga TTTCAGAAACGTGTTCCAGGACACAACAAGATGGAGTTTGAATCTTCACTGTATGAAGGA tcctttgaggaaatggatccacctgaaaatattgatgatatacaaagtgatctcatattc TCCTTTGAGGAAATGGATCCACCTGAAAATATTGATGATATACAAAGTGATCTCATATTC gtgaccctctctgtgaaggatagtaaaaygtctaccctctcctgtaagaacaagatcatt GTGACCCTCTCTGTGAAGGATAGTAAAAYGTCTACCCTCTCCTGTAAGAACAAGATCATT 240 ttcgttgacaaaagacagcctgtgttcgaggatatgactgatattgatcaaagtgccagt 120 gaaccccagaccagactgataatatacatgtacaaagacagtgaagtaagaggactggct 360 240 60 60 300

```
밁
Matches 471;
               Query Match
Best Local
                                                                                                                      The present sequence encodes a novel protein from mouse liver cells, which induces interferon-gamma (IFN gamma) production in immunocompetent cells. This protein enhances cytotoxicity of killer cells and induces their formation. It is used as an antioncotic agent for antitumour immunotherapy, an antiviral (including anti-AIDS) or antibacterial agent, and in the treatment of atopic or immune system diseases, e.g. asthma, hayfever or rheumatism. When formulated with interleukin-3, it is also used to treat leukopaenia and thrombocytopaenia associated with radiotherapy or chemotherapy of leukaemia and other cancers. When used in antitumour immunotherapy, this novel protein significantly improves the immunotherapeutic effect of interleukin-2 (IL-2), compared with use of IL-2 alone, either when administered to the patient (before administration of IL-2) or by addition to the medium in which cells
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Interferon-gamma, IFN-gamma; antiviral; antioncotic; radiotherapy; immunoregulatory; antitumour agent; chemotherapy; Leukopaenia; thrombocytopaenia; immunocompetent cell; asthma; hayfever; rheumatism; interleukin; killer cell; ds.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    T60536 standard; cDNA to mRNA; 471 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  EP767178-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   26-JAN-1998 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   26-SEP-1996;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    09-APR-1997
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   mat_peptide
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Mus musculus
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Mouse interferon-gamma inducer cDNA.
                                                                              Sequence 471 BP; 162 A; 91 C; 92 G; 125 T; 1 other;
                                                                                                              (intended for return to the patient) are being grown.
                                                                                                                                                                                                                                                                                                                                                                Disclosure; Page 22; 26pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                  tumours and as antimicrobial agent etc
                                                                                                                                                                                                                                                                                                                                                                                                               Human protein that induces interferon-gamma prodn. in immuno:competent cells - useful for adoptive immuno:therapy of
                                                                                                                                                                                                                                                                                                                                                                                                                                                                 WPI; 1997-205381/19.
P-PSDB; W15704.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                (HAYB ) HAYASHIBARA SEIBUTSU KAGAKU.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   20-SEP-1996;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ω
               Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Fujii M, Kurimoto M, Nukada Y,
Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                96JP-0269105.
95JP-0270725.
96JP-0067434.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   96EP-0306997
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   /product= interferon gamma inducer
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  /*tag=
               99.9%; Score 470.6; DB 18; 100.0%; Pred. No. 5.8e-124;
0; Mismatches
                              DB 18; Length 471;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Tanimoto T;
 Indels
 0;
 Gaps
 0;
```

```
밁
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     망
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Ş
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     밁
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  δÃ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Д
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      δÃ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Вb
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Qy
04-OCT-1995;
10-MAR-1995;
29-SEP-1995;
                                                                                                                                                                                                                                                                                                                                                                                                                                          Mus musculus
                                                                                                                                                                                                                                                                                                                                                                                                                                                                            viral disease;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Murine protein for induction of interferon-gamma
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           15-OCT-1997 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              T80210 standard; cDNA to mRNA; 471 BP
                               A drug containing a polypeptide which induces interferon-gamma useful for treating e.g. malignant tumours, viral, bacterial o
                                                                                                                                                                                                                                       24-JAN-1996;
                                                                                                                                                                                                                                                                         17-JUN-1997.
                                                                                                                                                                                                                                                                                                        JP09157180-A.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Interferon-gamma; immunocompetent cell; malignant tumour;
viral disease; bacterial infection; immune disease; ds.
                                                                                                                                (HAYB ) HAYASHIBARA SEIBUTSU KAGAKU
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 421 GAAAATGGGGATAAATCTGTAATGTTCACTCTCACTAACTTACATCAAAGT 471
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   181 gtgaccctctctgtgaaggatagtaaaaygtctaccctctcctgtaagaacaagatcatt 240
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     121 gaaccccagaccagactgataatatacatgtacaaagacagtgaagtaagaggactggct 180
                   immune diseases
                                                                                                   WPI; 1997-369391/34.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              421 gaaaatggggataaatctgtaatgttcactctcactaacttacatcaaagt 471
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                301 tttcagaaacgtgttccaggacacaacaagatggagtttgaatcttcactgtatgaagga 360
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                301 TTTCAGAAACGTGTTCCAGGACACAACAACAAGATGGAGTTTGAATCTTCACTGTATGAAGGA 360
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 241 tcctttgaggaaatggatccacctgaaaatattgatgatatacaaagtgatctcatattc 300
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      241 TCCTTTGAGGAAATGGATCCACCTGAAAATATTGATGATATACAAAGTGATCTCATATTC 300
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      181 GTGACCCTCTCTGTGAAGGATAGTAAAAYGTCTACCCTCTCCTGTAAGAACAAGATCATT 240
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      121 GAACCCCAGACCAGACTGATAATATACATGTACAAAGACAGTGAAGTAAGAGGACTGGCT 180
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                -PSDB; W24262.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       61 ttcgttgacaaaagacagcctgtgttcgaggatatgactgatattgatcaaagtgccagt 120
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          1 aactttggccgacttcactgtacaaccgcagtaatacggaatataaatgaccaagttctc 60
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           1 AACTTTGGCCGACTTCACTGTACAACCGCAGTAATACGGAATATAAATGACCAAGTTCTC 60
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             TTCGTTGACAAAAGACAGCCTGTGTTCGAGGATATGACTGATATTGATCAAAGTGCCAGT 120
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                95JP-0279906.
95JP-0078357.
95JP-0274988.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                            bacterial infection; immune
                                                                                                                                                                                                                                       96JP-0028722.
                                                                                                                                                                                                                                                                                                                                                                                          Location/Qualifiers
1..471
                                                                                                                                                                                                                                                                                                                                       /transl_except= pos:208..210, aa:Xaa
/note= "No stop codon given"
                                                                                                                                                                                                                                                                                                                                                                          /*tag= a
```

```
RESULT
V48227
                                                                                                                                                                                                                                                                                                                                                   밁
                                                                                                                                                                                                                                                                                                                                                                      Qy
                                                                                                                                                                                                                                                                                                                                                                                                   В
                                                                                                                                                                                                                                                                                                                                                                                                                       ρ
                                                                                                                                                                                                                                                                                                                                                                                                                                                      рь
                                                                                                                                                                                                                                                                                                                                                                                                                                                                             γ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Qy
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Qy
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          뫄
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   δÃ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        밁
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           밁
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               밁
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Qy
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 В
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Qy
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   SX CCCCCX B
                                                                                                                                                Mouse; interleukin-18; IL-18; osteoclast; hypercalcaemia; osteopenia; osteoclastoma Behcet's syndrome; osteosarcoma; arthropathy; osteoporos chronic rheumatoid arthritis; deformity ostitis; primary hyperthyroidi
                                                                                       Key
                        EP861663-A2.
                                                                                                                             Mus
                                                                                                                                                                                                       Mouse interleukin
                                                                                                                                                                                                                                16-NOV-1998
                                                                                                                                                                                                                                                           V48227;
                                                                                                                                                                                                                                                                                   V48227 standard;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   This sequence encodes a protein which induces interferon-gamma production in immuncompetent cells. This protein may be used as the major component in a drug for the prevention and treatment of e.g. malignant tumours, viral diseases, bacterial infections and
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Sequence 471 BP; 162 A;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Disclosure; Page
                                                                                                                                                                                                                                                                                                                                                   421
                                                                                                                                                                                                                                                                                                                                                                           421
                                                                                                                                                                                                                                                                                                                                                                                                    361
                                                                                                                                                                                                                                                                                                                                                                                                                                                        301
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  241
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         241
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           181
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     181
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             121
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     121
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                61
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        61
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  \vdash
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     1 AACTTTGGCCGACTTCACTGTACAACCGCAGTAATACGGAATATAAATGACCAAGTTCTC
                                                                                                                           sp.
                                                                                                                                                                                                                                                                                                             ഗ
                                                                                                                                                                                                                                                                                                                                                           GAAAATGGGGATAAATCTGTAATGTTCACTCTCACTAACTTACATCAAAGT 471
                                                                                                                                                                                                                                                                                                                                            gaaaatggggataaatctgtaatgttcactctcactaacttacatcaaagt 471
                                                                                                                                                                                                                                                                                                                                                                                                              TTTCAGAAACGTGTTCCAGGACACAACAAGATGGAGTTTGAATCTTCACTGTATGAAGGA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 TCCTTTGAGGAAATGGATCCACCTGAAAATATTGATGATATACAAAGTGATCTCATATTC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   GTGACCCTCTGTGAAGAGATAAAAAAAAGTCTACCCTCTCCTGTAAGAACAAGATCATT
                                                                                                                                                                                                                                                                                                                                                                                                tttcagaaacgtgttccaggacacaacaagatggagtttgaatcttcactgtatgaagga
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       gtgaccctctctgtgaaggatagtaaaaygtctaccctctcctgtaagaacaagatcatt
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          gaaccccagaccagactgataatatacatgtacaaagacagtgaagtaagaggactggct
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    GAACCCCAGACCAGACTGATAATATACATGTACAAAGACAGTGAAGTAAGAGGACTGGCT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          TTCGTTGACAAAAGACAGCCTGTGTTCGAGGATATGACTGATATTGATCAAAGTGCCAGT 120
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              aactttggccgacttcactgtacaaccgcagtaatacggaatataaatgaccaagttctc
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     tcctttgaggaaatggatccacctgaaaatattgatgatatacaaagtgatctcatattc
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ttcgttgacaaaagacagcctgtgttcgaggatatgactgatattgatcaaagtgccagt
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Conservative
                                                                                                                                                                                                                              (first entry)
                                            /rtag= a
/product= "Interleukin 18"
/note= "No stop or -"
                                                                                                   Location/Qualifiers
                                                                                                                                                                                                                                                                                   CDNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    10-11;
                                                                                                                                                                                                       18
                                                                                                                                                                                                       gene
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               99.9%;
                                                                                                                                                                                                                                                                                  ţ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                91 C;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 12pp; Japanese.
                                                                                                                                                                                                                                                                                  mRNA;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  0;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Score 470.6;
Pred. No. 5
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  92 G; 125
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Mismatches
                                                                                                                                                ceeccast; nypercatcaemia; osteopenia; ds;
osteosarcoma; arthropathy; osteoporosis;
formity ostitis; primary hyperthyroidism.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             5.8e-124;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               T; 1 other;
                                             codon given"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           DB 18;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Length
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 0;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  of
                                                                                                                                                                                                                                                                                                                                                                                                  420
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               300
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         180
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   180
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               60
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       60
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 0;
```

```
RESULT
Z36923
ID Z3
XX
AC Z3
XX
DT 13
                                                                                                                                                                     Вþ
                                                                                                                                                                                                                                              Qy
                                                                                                                                         δÃ
                                                                                                                                                                                           Qy
                                                                                                                                                                                                                       Ъ
                                                                                                                                                                                                                                                                         В
                                                                                                                                                                                                                                                                                               Qy
                                                                                                                                                                                                                                                                                                                           В
                                                                                                                                                                                                                                                                                                                                                  δÃ
                                                                                                                                                                                                                                                                                                                                                                              밁
                                                                                                                                                                                                                                                                                                                                                                                                    Qy
                                                                                                                                                                                                                                                                                                                                                                                                                                DЬ
                                                                                                                                                                                                                                                                                                                                                                                                                                                        γΩ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   В
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Qy
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Matches 470;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Query Match
Best Local
                          Z36923;
                                                   Z36923
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Interleukin-18 (IL-18) or a functional equivalent can be used for inhibition of osteoclast formation. IL-18 is used for treating or preventing osteoclast-related diseases e.g. hypercalcaemia, osteoclastoma Behcet's syndrome, osteosarcoma, arthropathy, chronic rheumatoid arthritis, deformity ostitis, primary hyperthyroidism, osteopenia and
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Use of interleukin-18 to inhibit osteoclast formation - of e.g. hypercalcaemia, osteoclastoma, Behcet's syndrome, osteosarcoma, chronic rheumatoid arthritis, deformity ost
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Sequence 471 BP; 162
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             osteoporosis.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Disclosure; Page 29; 56pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           primary hyperthyroidism and osteoporosis
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         WPI; 1998-448964/39
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Gillespie MT,
                                                                                                                  421
                                                                                                                                            421
                                                                                                                                                                     361
                                                                                                                                                                                               361
                                                                                                                                                                                                                       301
                                                                                                                                                                                                                                                 301
                                                                                                                                                                                                                                                                                                                           181
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             24-FEB-1998;
                                                                                                                                                                                                                                                                                                   241
                                                                                                                                                                                                                                                                                                                                                     181
                                                                                                                                                                                                                                                                                                                                                                              121
                                                                                                                                                                                                                                                                                                                                                                                                        121
                                                                                                                                                                                                                                                                                                                                                                                                                                 61
                                                                                                                                                                                                                                                                                                                                                                                                                                                        61
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              1 AACTTTGGCCGACTTCACTGTACAACCGCAGTAATACGGAATATAAATGACCAAGTTCTC
                                                                            σ
                                                                                                                                                                                                                   gaaaatggggataaatctgtaatgttcactctcactaacttacatcaaagt
                                                                                                                                            GAAAATGGGGGATAAATCTGTAATGTTCACTCTCACTAACTTACATCAAAGT
                                                                                                                                                                               GTGACCCTCTCTGTGAAGGATAGTAAAAYGTCTACCCTCTCCTGTAAGAACAAGATCATT 240
                                                                                                                                                                                                                                                                                                                                                                                          GAACCCCAGACCAGACTGATAATATACATGTACAAAGACAGTGAAGTAAGAGGACTGGCT 180
                                                                                                                                                                                                                                                                                                                                                                                                                                                                               aactttggccgacttcactgtacaaccgcagtaatacggaatataaatgaccaagttctc 60
                                                                                                                                                                   cactttcttgcttgccaaaaggaagatgatgctttcaaactcattctgaaaaaaaggat
                                                                                                                                                                                                                                                                     tcctttgaggaaatggatccacctgaaaatattgatgatatacaaagtgatctcatattc
                                                                                                                                                                                                                                                                                                TCCTTTGAGGAAATGGATCCACCTGAAAATATTGATGATATACAAAGTGATCTCATATTC 300
                                                                                                                                                                                                                                                                                                                        gtgaccctctctgtgaaggatagtaaaatgtctaccctctcctgtaagaacaagatcatt
                                                                                                                                                                                                                                                                                                                                                                           gaaccccagaccagactgataatatacatgtacaaagacagtgaagtaagaggactggct
                                                                                                                                                                                                                                                                                                                                                                                                                             ttcgttgacaaaagacagcctgtgttcgaggatatgactgatattgatcaaagtgccagt 120
                                                                                                                                                                                                                                                                                                                                                                                                                                         TTCGTTGACAAAAGACAGCCTGTGTTCGAGGATATGACTGATATTGATCAAAGTGCCAGT 120
                                                                                                                                                                                                                                                                                    Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Horwood NJ,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    97JP-0055468
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             98EP-0301352
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               99.98;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   2
                                                   ç
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  91 C;
                                                   mRNA;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Kurimoto M,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Score 470.6; DB 19
Pred. No. 5.8e-124;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           KAGAKU
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   92 G;
                                                    471
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  126
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Τ,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Udagawa
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  0 other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            19;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Length
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ostitis

    in treatment

                                                                                                                  471
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    0;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Gaps
                                                                                                                                                                                                                                              360
                                                                                                                                                                                                                    360
                                                                                                                                                                                                                                                                                                                                                                           180
                                                                                                                                                                                                                                                                        300
                                                                                                                                                                                                                                                                                                                        240
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          60
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  0
```

13-MAR-2000

(first entry)

```
Qγ
                                                                                                                                                                  밁
                                                                                                                                                                                                            The present sequence encodes a murine protein that induces interferon (IFN) gamma production by immunocompetent cells. IFN-gamma is a protein which has antiviral, antioncotic and immunoregulatory activities, and is produced by immunocompetent cells stimulated with antigens or mitogens. A probe derived from the present sequence was used to isolate the corresponding human protein from human liver cells. The protein of the invention is used to treat IFN-gamma susceptible diseases, and also have use as a antiviral agent, antibacterial agent, antitumour agent, compared to the careful agent and blood platelet enhancing agent. Diseases which can be treated with the protein include viral diseases such as Combined tumours such as and chronic granulomatous disease; blood call malignant tumours such as renal call gament tumours such as renal call gament such as adult T cell leukaemia, and immune diseases
                                                                         Matches 471;
                                                                                                         Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          10-MAR-1995;
18-SEP-1995;
29-SEP-1995;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Mouse; interferon gamma production; IFN-gamma; immunocompetent cell; antiviral; immunoregulatory; antigen; mitogen; IFN-gamma susceptible disease; antibacterial; antitumour; IFN-gamma susceptible disease; antibacterial; antitumour; blood platelet enhancing agent; hepatitis; herpes syndrome; condyloma; blood platelet enhancing agent; hepatitis; herpes syndrome; condyloma; blood platelet enhancing agent; hepatitis; solid malignant tumour; bronic granulomatous disease; renal cancer; mycosis fungoides; bronic granulomatous disease; blood cell malignant tumour; adult T cell leukaemia;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         DNA encoding a protein that induces IFN-gamma production
                                                                                                                                                                Sequence 471 BP; 162 A; 91 C; 92 G; 125 T; 1 other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Disclosure; Page 3; 42pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 (HAYB ) HAYASHIBARA SEIBUTSU KAGAKU.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         EP962531-A2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Mus sp.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         chronic myelogenous leukaemia; malignant leukaemia; immune disease;
                                                                                                                                                                                                  such as allergy and rheumatism.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   susceptible diseases
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Novel polypeptides used in the treatment of interferon-gamma
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    10-NOV-1995;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   10-NOV-1995;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   mat_peptide
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          allergy; rheumatism; ds.
                                                                                            Loca!
1 aactttggccgacttcactgtacaaccgcagtaatacggaatataaaatgaccaagttctc 60
                                   1 AACTTTGGCCGACTTCACTGTACAACCGCAGTAATACGGAATATAAATGACCAAGTTCTC 60
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           2000-064289/06
                                                                                          Similarity
                                                                       Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       95JP-0262062.
95JP-0274988.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          94JP-0304203.
95JP-0058240.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    95EP-0308055.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           95JP-0078357
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 99EP-0104104.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          /transl_except= (pos: 208..210, aa: Xaa)
/note= "Xaa is not specified"
                                        99.9%; ...
100.0%; Pred. No.
170 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Tanimoto T, Okamura H;
                                                                                        Score 470.6; DB 21; Pred. No. 5.8e-124;
                                                                       Indels
                                                                                                         Length 471;
                                                                       0;
                                                                       Gaps
                                                                       0;
```

```
δõ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Š
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              밁
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Ъ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Q
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 밁
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Вþ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Ωy
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Дb
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Дb
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        QY
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       DЬ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Qy
 Interferon-gamma inducing factor; interferon-gamma; killer cell; antitumour agent; antiviral agent; antimicrobial agent; tumour; mIGIF; hepatitis; malaria; tuberculosis; renal carcinoma; rheumatism; AIDS; osteoporosis; thrombopenia; acquired immunodeficiency syndrome; ds.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        V32755;
(HAYB ) HAYASHIBARA SEIBUTSU KAGAKU
                                                                                                                                                                                                                                                                                  sig_peptide
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   V32755 standard; cDNA; 570 BP
                             21-JAN-1997;
                                            14-NOV-1997;
29-NOV-1996;
                                                                                                                     03-JUN-1998
                                                                                                                                                EP845530-A2
                                                                                                                                                                                                                       mat_peptide
                                                                                                                                                                                                                                                                                                                                                          5'UTR
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Wild-type mouse interferon-gamma inducing factor cDNA.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          25-SEP-1998
                                                                                        28-NOV-1997;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          421 gaaaatggggataaatctgtaatgttcactctcactaacttacatcaaagt 471
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     301 tttcagaaacgtgttccaggacacaacaagatggagtttgaatcttcactgtatgaagga
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              301
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   181 gtgaccctctctgtgaaggatagtaaaaygtctaccctctcctgtaagaacaagatcatt
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  181
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         121 GAACCCCAGACCAGACTGATAATATACATGTACAAAGACAGTGAAGTAAGAGGACTGGCT 180
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       421 GAAAATGGGGATAAATCTGTAATGTTCACTCTCACTAACTTACATCAAAGT 471
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          241 tcctttgaggaaatggatccacctgaaaatattgatgatatacaaagtgatctcatattc
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       241 TCCTTTGAGGAAATGGATCCACCTGAAAATATTGATGATATACAAAGTGATCTCATATTC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      61 TTCGTTGACAAAAGACAGCCTGTGTTCGAGGATATGACTGATATTGATCAAAGTGCCAGT 120
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       61 ttcgttgacaaaagacagcctgtgttcgaggatatgactgatattgatcaaagtgccagt 120
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             TTTCAGAAACGTGTTCCAGGACACAACAAGATGGAGTTTGAATCTTCACTGTATGAAGGA 360
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    GTGACCCTCTCTGTGAAGGATAGTAAAAYGTCTACCCTCTCTCTGTAAGAACAAGATCATT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              gaaccccagaccagactgataatatacatgtacaaagacagtgaagtaagaggactggct 180
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   (first entry)
                                            97JP-0329715
96JP-0333037
                               97JP-0020906
                                                                                         97EP-0309632
                                                                                                                                                                                          /*tag= d
559..570
                                                                                                                                                                                                                                                                                  /product= "Immature mouse IGIF" 16..84
                                                                                                                                                                                                                                                                                                                             /*tag=
16..558
                                                                                                                                                                                                                       85..555
                                                                                                                                                                                                                                   /note= "This sequence claimed by the inventors under claim 11 in the specification"
                                                                                                                                                                                                                                                                                                                                                                        Location/Qualifiers
                                                                                                                                                                                                           Q
```

```
RESULT
V32633
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             뫄
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Matches 470;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Query Match
Best Local
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          the consensus sequences shown in W48956-W48958. The mutant mIGIFs are capable of stimulating immunocompetent cells for the production of interferon-gamma and are claimed to be less toxic, more active and stable than the corresponding wild type interferon-gamma inducing factor. The mutant mIGIFs are also claimed to enhance killer cell cytotoxicity and/or induce killer cell formation, and may therefore be useful as antitumour agents, antitumour immunotherapeutics, antiviral agents and antimicrobial agents. The mutant mIGIFs are also claimed to be useful for treating hepatitis, acquired immunodeficiency syndrome (AIDS), malaria, tuberculosis, solid malignant tumours (e.g. renal carcinoma), rheumatism, osteoporosis and thrombopenia caused by radiation- and observed.
V32633;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 The present sequence represents the wild-type mouse interferon-gamma inducing factor (mIGIF) cDNA. The invention provides for mutant mouse and human interferon-gamma inducing factors in which one or more cysteine residues are replaced with different residues at or away from the consensus sequences shown in W48956-W48958. The mutant mIGIFs are
                                                V32633 standard;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Sequence 570 BP; 175 A; 123 C; 121 G;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Claim 11; pages 38-39; 59pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Mutants of interferon-gamma inducing polypeptide - useful antitumour, antiviral, antimicrobial or anti-immunopathic
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         carcinoma), rheumatism, osteopradiation- and chemo-therapy
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Kurimoto M,
                                                                                                                                                                                                                                                                                                          445
                                                                                                                                                                                                                                                                                                                                                                   361
                                                                                                                                                                                                                                                                                                                                                                                                                                                                           301
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 325
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           265
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               181
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                145
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   61
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             85
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              1 AACTTTGGCCGACTTCACTGTACAACCGCAGTAATACGGAATATAAATGACCAAGTTCTC
                                                                                                                                                                                                                    GAAAATGGGGATAAATCTGTAATGTTCACTCTCACTAACTTACATCAAAGT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               GAACCCCAGACCAGACTGATAATATACATGTACAAAGACAGTGAAGTAAGAGGACTGGCT
                                                                                                                                                                                       gaaaatggggataaatctgtaatgttcactctcactaacttacatcaaagt
                                                                                                                                                                                                                                                                                                    cactttcttgcttgccaaaaggaagatgatgctttcaaactcattctgaaaaaaaggat
                                                                                                                                                                                                                                                                                                                                                                 TCCTTTGAGGAAATGGATCCACCTGAAAATATTGATGATATACAAAGTGATCTCATATTC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  GTGACCCTCTCTGTGAAGGATAGTAAAAXGTCTACCCTCTCCTGTAAGAACAAGATCATT
                                                                                                                                                                                                                                                                                                                                                                                                                    tttcagaaacgtgttccaggacacaacaagatggagtttgaatcttcactgtatgaagga
                                                                                                                                                                                                                                                                                                                                                                                                                                                                     TTTCAGAAACGTGTTCCAGGACACAACAAGATGGAGTTTGAATCTTCACTGTATGAAGGA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         gtgaccctctctgtgaaggatagtaaaatgtctaccctctcctgtaagaacaagatcatt
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                gaaccccagaccagactgataatatacatgtacaaagacagtgaagtaagaggactggct
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ttcgttgacaaaagacagcctgtgttcgaggatatgactgatattgatcaaagtgccagt
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   TTCGTTGACAAAAGACAGCCTGTGTTCGAGGATATGACTGATATTGATCAAAGTGCCAGT 120
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       aactttggccgacttcactgtacaaccgcagtaatacggaatataaatgaccaagttctc 144
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           tcctttgaggaaatggatccacctgaaaatattgatgatatacaaagtgatctcatattc
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        1998-288747/26
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 W48960.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Okamoto
                                                   cDNA; 471 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              99.98;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Τ,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Yamamoto
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Score 470.6; DB 19
Pred. No. 6.2e-124;
1; Mismatches 0;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           151
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Τ;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           DB 19;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           0 other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       0;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Length 570;
                                                                                                                                                                                                                                                 471
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           agents
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       as
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    0;
                                                                                                                                                                                                                                                                                                                                                                                                                    444
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      180
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             204
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              384
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   264
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    0,
```

₹X H

밁 Q

Matches 469;

Similarity

99.6%;

Score 469; DB 19; Pred. No. 1.6e-123; Mismatches

Length 471;

Indels

0

Gaps

Conservative

ب ۳

AACTTTGGCCGACTTCACTGTACAACCGCAGTAATACGGAATATAAATGACCAAGTTCTC 60 aactttggccgacttcactgtacaaccgcagtaatacggaatataaatgaccaagttctc

Ωy В ρ

맑

Qy 밁 δÃ

밁

γQ Вb

밁

ф

δÃ

Qy

Qy

```
for mutant human and mouse interferon gamma inducing factors in which one or more cysteine residues are replaced with different residues at or away from the consensus sequences shown in w48956-W48958. The mutant mIGIFs are capable of stimulating immunocompetent cells for the production of interferon-gamma and are claimed to be less toxic, more active and stable than the corresponding wild type mIGIF factor. The mutant mIGIFs are also claimed to enhance killer cell cytotoxicity and/or induce killer cell formation, and may therefore be useful as antitumour agents. The mutant mIGIFs are also claimed to be useful as antitumour agents. The mutant mIGIFs are also claimed to be useful for treating hepatitis, acquired immunodeficiency syndrome (AIDS), malaria, tuberculosis, solid malignant tumours (e.g. renal carcinoma), rheumatism, osteoporosis and thrombopenia caused by radiation- and chemo-therapy.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   14-NOV-1997;
29-NOV-1996;
21-JAN-1997;
                                                                                                                                                                                                                                                                                                                                       Mutants of interferon-gamma inducing polypeptide - useful as antitumour, antiviral, antimicrobial or anti-immunopathic agents
Sequence 471 BP; 163 A; 91 C; 92 G;
                                                                                                                                                                                                                                                       The present sequence represents the mutant mouse inducing factor cDNA mIGIF/MUT12. The wild-type
                                                                                                                                                                                                                                                                                                         Claim 10;
                                                                                                                                                                                                                                                                                                                                                                                                       WPI; 1998-288747/26.
                                                                                                                                                                                                                                                                                                                                                                                                                                       Kurimoto M,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  03-JUN-1998
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                mutation
                                                                                                                                                                                                                                          inducing factor cDNA mIGIF/MUT12. The wild-type mouse interferon-gamma factor (mIGIF) cDNA sequence is shown in V32755. The invention provide:
                                                                                                                                                                                                                                                                                                                                                                                        P-PSDB; W48969
                                                                                                                                                                                                                                                                                                                                                                                                                                                                     (HAYB ) HAYASHIBARA SEIBUTSU KAGAKU.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  EP845530-A2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Synthetic
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Mus sp.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           osteoporosis;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        hepatitis;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           antitumour
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  28-NOV-1997;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Interferon-gamma inducing factor; interferon-gamma; killer
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Mutant mouse interferon-gamma inducing factor cDNA mIGIF/MUT12
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         25-SEP-1998
                                                                                                                                                                                                                                                                                                       page 50; 59pp;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        agent; antiviral agent; antimicrobial agent; tumour; mIGI
malaria; tuberculosis; renal carcinoma; rheumatism; AIDS;
is; thrombopenia; acquired immunodeficiency syndrome; ds.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                     Okamoto
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   97JP-0329715.
96JP-0333037.
97JP-0020906.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  97EP-0309632
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             /note= "CDS of 373..375
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          /*tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              /*tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                       Į,
                                                                                                                                                                                                                                                                                                       English
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              does not contain a
125 T; 0 other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            stop
                                                                                                                                                                                                                                      The invention provides
                                                                                                                                                                                                                                                                           interferon-gamma
```

```
멍
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 QΥ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   DЬ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                멍
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Qy
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        δÃ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Qy
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Qy
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            В
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Дb
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    δÃ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Db
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               20
            Kurimoto M,
                                                                                     14-NOV-1997;
29-NOV-1996;
                                                                                                                                                                                                                                                                           mutation
                                          (HAYB ) HAYASHIBARA SEIBUTSU KAGAKU.
                                                                                                                                                                  03-JUN-1998.
                                                                                                                                                                                              EP845530-A2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Mutant mouse interferon-gamma inducing factor cDNA mIGIF/MUT11.
                                                                         21-JAN-1997;
                                                                                                                                   28-NOV-1997;
                                                                                                                                                                                                                                                                                                                                                                                                  Synthetic.
                                                                                                                                                                                                                                                                                                                                                                                                                                             osteoporosis; thrombopenia; acquired immunodeficiency syndrome; ds
                                                                                                                                                                                                                                                                                                                                                                                                                                                           antitumour agent; antiviral agent; antimicrobial agent; tumour; mIGI hepatitis; malaria; tuberculosis; renal carcinoma; rheumatism; AIDS;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Interferon-gamma inducing factor; interferon-gamma; killer cell;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      25-SEP-1998 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   V32632 standard; cDNA; 471 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          421 GAAAATGGGGATAAATCTGTAATGTTCACTCTCACTAACTTACATCAAAGT 471
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   301
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        421 gaaaatggggataaatctgtaatgttcactctcactaacttacatcaaagt 471
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   361 cactttcttgctagccaaaaggaagatgatgctttcaaactcattctgaaaaaaaggat
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     301
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    121 GAACCCCAGACCAGACTGATAATATACATGTACAAAGACAGTGAAGTAAGAGGACTGGCT 180
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  61 TTCGTTGACAAAAGACAGCCTGTGTTCGAGGATATGACTGATATTGATCAAAGTGCCAGT 120
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                tttcagaaacgtgttccaggacacaacaagatggagtttgaatcttcactgtatgaagga
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                TTTCAGAAACGTGTTCCAGGACACAACAAGATGGAGTTTGAATCTTCACTGTATGAAGGA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            tcctttgaggaaatggatccacctgaaaatattgatgatatacaaagtgatctcatattc
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           TCCTTTGAGGAAATGGATCCACCTGAAAATATTGATGATATACAAAGTGATCTCATATTC 300
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         gtgaccctctctgtgaaggatagtaaaatgtctaccctctcctgtaagaacaagatcatt 240
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       GTGACCCTCTCTGTGAAGGATAGTAAAAYGTCTACCCTCTCCTGTAAGAACAAGATCATT 240
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    gaaccccagaccagactgataatatacatgtacaaagacagtgaagtaagaggactggct 180
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ttcgttgacaaaagacagcctgtgttcgaggatatgactgatattgatcaaagtgccagt 120
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Okamoto I,
                                                                                     97JP-0329715.
96JP-0333037.
                                                                         97JP-0020906
                                                                                                                                   97EP-0309632
                                                                                                                                                                                                                                                                       /note= "CDS does not contain a stop codon" 19..21
                                                                                                                                                                                                                                       /note= "changed from TGT in wild-type to GCT in
                                                                                                                                                                                                                                                                                                                  /product= "Mutant human interferon-gamma inducing
                                                                                                                                                                                                                                                                                                                                                                Location/Qualifiers
                                                                                                                                                                                                                           mutant"
            Yamamoto K;
                                                                                                                                                                                                                                                                                                       factor mIGIF/MUT11"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                300
```

RESULT 10

V20875 standard; cDNA;

28-JUL-1998 (first entry)

Вþ

421 GAAAATGGGGATAAATCTGTAATGTTCACTCTCACTAACTTACATCAAAGT 471

421 gaaaatggggataaatctgtaatgttcactctcactaacttacatcaaagt 471

```
Вb
                                                                                                               B
                                                                                                                                                               δÃ
                                                                                                                                                                                                                           Вb
                                                                                                                                                                                                                                                                           ρy
                                                                                                                                                                                                                                                                                                                                        Дb
                                                                                                                                                                                                                                                                                                                                                                                        δÃ
                                                      Qy
                                                                                                                                                                                                                                                                                                                                                                                                                                                    Вþ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    QΥ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Вþ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Qy
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         SXXCCCCCCCCCCCCCXXX
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               The present sequence represents the mutant mouse interferon-gamma conducting factor cDNA mIGIF/MUTI1. The wild-type mouse interferon-gamma conducting factor cDNA mIGIF/MUTI1. The wild-type mouse interferon-gamma conduction (mIGIF) cDNA sequence is shown in V32755. The invention provides conforme cysteine residues are replaced with different residues at or away conformed the consensus sequences shown in W48956-W48958. The mutant mIGIFs care capable of stimulating immunocompetent cells for the production of conterferon-gamma and are claimed to be less toxic, more active and conformation, and may therefore be useful as antitumour agents, are also claimed to enhance killer cell cytotoxicity and/or induce killer cell formation, and may therefore be useful as antitumour agents. The mutant mIGIFs are also claimed to be useful as antitumour agents. The mutant mIGIFs are also claimed to be useful for treating hepatitis, careful immunodeficiency syndrome (AIDS), malaria, tuberculosis, solid malignant tumours (e.g. renal carcinoma), rheumatism, osteoporosis and continuours caused by radiation— and chemo-therapy.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Sequence 471 BP; 162 A; 92 C; 92 G; 125 T; 0 other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Claim 10; pages 49-50; 59pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Mutants of interferon-gamma inducing polypeptide - useful as antitumour, antiviral, antimicrobial or anti-immunopathic agents
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   WPI; 1998-288747/26.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                P-PSDB; W48968.
301 tttcagaaacgtgttccaggacacaacaagatggagtttgaatcttcactgtatgaagga 360
                                                      301 TTTCAGAAACGTGTTCCAGGACACAACAAGATGGAGTTTGAATCTTCACTGTATGAAGGA 360
                                                                                                         241 tcctttgaggaaatggatccacctgaaaatattgatgatatacaaagtgatctcatattc 300
                                                                                                                                  241 TCCTTTGAGGAAATGGATCCACCTGAAAATATTGATGATATACAAAGTGATCTCATATTC 300
                                                                                                                                                                                                                     181 gtgaccctctctgtgaaggatagtaaaatgtctaccctctcctgtaagaacaagatcatt 240
                                                                                                                                                                                                                                                                           181 GTGACCCTCTCTGTGAAGGATAGTAAAAYGTCTACCCTCTCCTGTAAGAACAAGATCATT 240
                                                                                                                                                                                                                                                                                                                              121 gaaccccagaccagactgataatatacatgtacaaagacagtgaagtaagaggactggct 180
                                                                                                                                                                                                                                                                                                                                                                                        121 GAACCCCAGACCAGACTGATAATATACATGTACAAAGACAGTGAAGTAAGAGGACTGGCT 180
                                                                                                                                                                                                                                                                                                                                                                                                                                           61 ttcgttgacaaaagacagcctgtgttcgaggatatgactgatattgatcaaagtgccagt 120
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    61 TTCGTTGACAAAAGACAGCCTGTGTTCGAGGATATGACTGATATTGATCAAAGTGCCAGT 120
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    1 aactttggccgacttcacgctacaaccgcagtaatacggaatataaatgaccaagttctc 60
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Match 99.2%;
Local Similarity 99.4%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             1 AACTTTGGCCGACTTCACTGTACAACCGCAGTAATACGGAATATAAATGACCAAGTTCTC 60
                                                                                                                                                                                                                                                  Conservative

    Mismatches

                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Score 467.4; DB 19; Length 471; Pred. No. 4.6e-123;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   0;
```

0;

```
а
                                                                                                                                                                                                                                                                                                                                                             В
                                                                                                                                                                                                                                                                                                                                                                                                   QΥ
                                                                                                                                                                                                                                                                                                                                                                                                                                                        밁
                                                                                   뮰
                                                                                                                                                                                밁
밁
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Query Match
Best Local
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 This is the nucleotide sequence of the rat interferon-gamma inducing factor (IGIF), also known as interleukin-18 (IL-18). It can be used to transform a cell, which upon its expression can cause the cell to produce rat IGIF, i.e. IL-18 or IL-18 alpha. The antibody to IGIF, IGIF and probes derived from it, are useful for detection of IL-18 or IL-18 alpha present in a sample. The amount of IL-18 or IL-18 alpha in a sample can be used to quantitate stress in a mammal.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 CDS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Sequence 722 BP; 252 A; 138 C; 140 G;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Claim 4; Page 31; 47pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Rat interferon-gamma inducing factors and related DNA - useful for quantitating stress in a mammal % \left( 1\right) =\left\{ 1\right\} 
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Conti B,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 08-APR-1997;
09-SEP-1996;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        08-SEP-1997;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   12-MAR-1998
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              WO9810072-AI
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              sig_peptide
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Rattus
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Rat interferon-gamma inducing factor; IGIF; interleukin-18; IL-18;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Nucleotide sequence encoding a rat interferon-gamma inducing factor
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      (CORR ) CORNELL RES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            IL-18-alpha;
410
                                                                                                                                                                                                                             179
                                                                                                                                                                                                                                                                          230
                                                                                                                                                                                                                                                                                                                                                                                                                                                          110
                                                                                                                                                                                                                                                                                                                                                                                                             62
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                             2 ACTITGGCCGACTICACTGTACAACCGCAGTAATACGGAATATAAATGACCAAGTTCTCT 61
                                                                                                                                                                                                                                                                                                                                                                                        TCGTTGACAAAAGA----CAGCCTGTGTTCGAGGATATGACTGATATTGATCAAAGTGCCA 118
                                                                                                                                                                                                                                                                                                                                                                                                                                                   actttggcagacttcactgtacaaccgcagtaatacggagcataaatgaccaagttctct 169
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              1998-193622/17
tctttcagaaacgtgtgccaggacacaacaaaatggaatttgaatcttccctgtatgaag
                   TCTTTCAGAAACGTGTTCCAGGACACAACAAGATGGAGTTTGAATCTTCACTGTATGAAG 358
                                                                                                                                                                                                       CTGTGACCCTCTCTGTGAAGGATAGTAAAAYGTCTACCCTCTCTCTGTAAGAACAAGATCA 238
                                                                                                                                                                                                                                                                                                                       GTGAACCCCAGACCAGACTGATAATATACATGTACAAAGACAGTGAAGTAAGAGGACTGG 178
                                                                                     tttcctttgaggaaatgaatccacctgaaaatattgatgatataaaaagtgatctcatat 409
                                                                                                             TTTCCTTTGAGGAAATGGATCCACCTGAAAATATTGATGATATACAAAGTGATCTCATAT
                                                                                                                                                                           ctgtgaccctatctgtgaaggatggaaggatgtctaccctctcctgtaaaaacaaaatca
                                                                                                                                                                                                                                                                          acgaatcccagaccagactgataatatatgtacaaagatagtgaagtaagaggactgg
                                                                                                                                                                                                                                                                                                                                                                tcgttgacaaaagaaacccgcctgtgttcgaggacatgcctgatatcgaccgaacagcca
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                434;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   qs
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        W53282.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Joh TH;
                                                                                                                                                                                                                                                                                             Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              transformation; antibody; probe;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 97US-0043087.
96US-0025141.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      97WO-US15891
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    /product= "interferon-gamma inducing
1..108
/*tag= b
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           /*tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        FOUND INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     85.1%;
91.8%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Score 400.6; DB 19
Pred. No. 4.1e-104;
1; Mismatches 35;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                192 T; 0 other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           DB 19;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              hybridisation;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Length 722;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        factor"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ω,
  469
                                                                                                                                                                                    349
                                                                                                                                                                                                                                                                          289
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ۲,
```

δδ

δÃ

Q

δÃ

γQ 밁 Ş

TCGTTGACAAAAGA----CAGCCTGTGTTCGAGGATATGACTGATATTGATCAAAAGTGCCA 118 actttggcagacttcactgtacaaccgcagtaatacggagcataaatgaccaagttctct 169 ACTTTGGCCGACTTCACTGTACAACCGCAGTAATACGGAATATAAATGACCAAGTTCTCT 61

110

N

Matches

Similarity

Conservative

<u>,</u>

Mismatches

ω ;

Indels

60;

Gaps

Ņ,

Qy

```
RESULT
V20876
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  멍
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Qy
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 В
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Q
                                                                      This is the nucleotide sequence of the rat interferon-gamma inducing factor (IGIF) isoform, also known as interleukin-18-alpha (IL-18-alpha). It can be used to transform a cell, which upon its expression can cause the cell to produce rat IGIF, i.e. IL-18 or IL-18 alpha. It is made by the deletion of 57 bases (360-417) from 720875, a probable exon. The antibody to IGIF, IGIF and probes derived from it, are useful for detection of IL-18 or IL-18 alpha present in a sample. The amount of IL-18 or IL-18 alpha in a sample can be used to quantitate stress in a mammal.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            CDS
                                                                                                                                                                                                                      Claim 7; Pages 32-333; 47pp; English.
                                                                                                                                                                                                                                            quantitating stress in a mammal
                                                                                                                                                                                                                                                                                                                      Conti B,
                                                                                                                                                                                                                                                                                                                                                                        09-SEP-1996;
                                                                                                                                                                                                                                                                                                                                                                                                                                                            WO9810072-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Rattus
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Rat interferon-gamma inducing factor; IGIF; interleukin-18;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Nucleotide sequence encoding a rat interleukin-18-alpha
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     28-JUL-1998
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              V20876;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    V20876 standard; cDNA; 665
                                                                     Sequence 665 BP; 229 A; 130 C; 132 G; 174 T; 0 other;
                                                                                                                                                                                                                                                         Rat interferon-gamma inducing factors and
                                                                                                                                                                                                                                                                                  P-PSDB; W53283
                                                                                                                                                                                                                                                                                               WPI; 1998-193622/17
                                                                                                                                                                                                                                                                                                                                                                                    08-APR-1997;
                                                                                                                                                                                                                                                                                                                                                                                                             08-SEP-1997;
                                                                                                                                                                                                                                                                                                                                                                                                                                    12-MAR-1998
                                                                                                                                                                                                                                                                                                                                              (CORR ) CORNELL RES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       IL-18-alpha; transformation; antibody; probe; hybridisation;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 530 atgaaaatggggataaatctgtaatgttcactcttactaacttacatcaaagt 582
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              419 ATGAAAATGGGGATAAATCTGTAATGTTCACTCTCACTAACTTACATCAAAGT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Local 379;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             11
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 sp.
                                                                                                                                                                                                                                                                                                                        Joh
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     (first entry)
                                                                                                                                                                                                                                                                                                                                                                        97US-0043087.
96US-0025141.
                                                                                                                                                                                                                                                                                                                                                                                                             97WO-US15891
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Location/Qualifiers
1..528
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   /product= "interleukin-18-alpha"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 /*tag=
                                                                                                                                                                                                                                                                                                                                                 FOUND INC
                      59.4%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ВP
                      Score 279.8; DB 1
Pred. No. 5.5e-70;
                                                                                                                                                                                                                                                            related
                                 DB 19;
                                                                                                                                                                                                                                                             DNA
                                   Length 665;

    useful for

                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      529
```

```
Qy
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                DЬ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Qy
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Вþ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      γQ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        В
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Qy
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Ъ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                δÃ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 В
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             QΥ
     255624;
The cDNA was produced from alveolar macrophage mRNA via reverse transcription using primer 255625, and the CDNA amplified via PCR using primers 255626-255629. The invention relates to the use of recombinant IL-18 as a vaccine adjuvant. Adjuvants are used in vaccines to potentiate the immune response to an antigen derived from the pathogen. It is important that the correct type of immune reaction is triggered,
                                                                                                                                                                                                                                                                                                                                                                                          W09956775-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Interleukin-18; IL-18; adjuvant; vaccine; immune reaction;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Equine interleukin-18 (IL-18) cDNA.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               27-MAR-2000 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Z55624 standard; cDNA; 582 BP.
                                                                                                                         Claim 11; Page 22; 28pp; English.
                                                                                                                                                                                                                                                                                                07-MAY-1998;
                                                                                                                                                                                                                                                                                                                               04-MAY-1999;
                                                                                                                                                                                                                                                                                                                                                             11-NOV-1999.
                                                                                                                                                                                                                                     Nicolson L,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    419 ATGAAAATGGGGATAAATCTGTAATGTTCACTCACTAACTTACATCAAAGT 471
                                                                                              This sequence represents cDNA encoding equine interleukin-18 (IL-18).
                                                                                                                                                        Novel vaccine adjuvant used to increase the immune response
                                                                                                                                                                                                                                                                   (ALKU ) AKZO NOBEL NV
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   473 atgaaaatggggataaatctgtaatgttcactcttactaacttacatcaaagt 525
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             359 -----agaaacgtgtgccaggacacaacaacaatggaatttgaatcttccctgtatgaag 412
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       350 tttcctttg-----
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      179 CTGTGACCCTCTCTGTGAAGGATAGTAAAAYGTCTACCCTCTCCTGTAAGAACAAGATCA 238
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 230 acgaatcccagaccagactgataatatatgtacaaagatagtgaagtaagaggactgg
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               119 GTGAACCCCAGACCAGACTGATAATATACATGTACAAAGACAGTGAAGTAAGAGGACTGG 178
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            299 TCTTTCAGAAACGTGTTCCAGGACACAACAAGATGGAGTTTGAATCTTCACTGTATGAAG 358
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       239 TTTCCTTTGAGGAAATGGATCCACCTGAAAATATTGATGATATACAAAGTGATCTCATAT 298
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    290 ctgtgaccctatctgtgaaggatggaaggatgtctaccctctcctgtaaaaacaaaatca 349
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                2000-072212/06.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Y58241
                                                                                                                                                                                                                                     Rijke EO;
                                                                                                                                                                                                                                                                                                  98EP-0201451.
                                                                                                                                                                                                                                                                                                                               99WO-EP03098.
                                                                                                                                                                                                                                                                                                                                                                                                                        /product= "Equine IL-18"
                                                                                                                                                                                                                                                                                                                                                                                                                                     /*tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    289
```

```
888888888888888888
                                               CC since many types of immune mechanisms that can be activated are CC inadequate for the control of a particular pathogen. Mice were injected CC intramuscularly with a vaccine formulation either containing inactivated CC seedorables virus (PRV) plus tetanus toxoid (PT), or a formulation CC containing inactivated PRV, TT and 0.1 micrograms of recombinant murine CC II.18. When subsequently challenged with virulent PRV, unvaccinated CC control mice all succumbed to the infection, and only 30% of mice CC vaccinated with vaccine antigen alone (inactivated PRV plus TT) survived CC the infection. In contrast, mice which received the same amount of CC vaccine antigen in conjunction with II.18 had an 80% survival rate CC immunisation in conjunction with II.18 had an 80% survival rate CC immunisation funans and other animals, such as plys, sheep, birds, CC cattle, dogs, horses and fish. An adjuvant composition comprising CC II.18 may be administered concomitantly or sequentially with a vaccine CC transcriptional regulatory sequences may be used in DNA vaccines for the CC in vivo expression of II.18 in the cells of a vaccinated animal.
Sequence 582 BP; 200 A; 102 C; 113 G; 167 T; 0 other;
```

Ņ

δÃ QY 8 δÃ 뫄 Ωy 밁 B В Q Д Ω δÃ Matches 355; Query Match 179 CTGTGACCCTCTCTGTGAAGGATAGTAAAAYGTCTACCCTCTCCTGTAAGAACAAGATCA 238 110 actttggcaggcttgaacctaaactctcaatcatacgaaaatttgaacgaccaagttctct 169 530 aggatgaaaatggggataaatctgtaatgttcactgttcaaaac 573 416 AGGATGAAAATGGGGGATAAATCTGTAATGTTCACTCTCACTAAC 459 356 AAGGACACTTTCTTGCCTTGCCAAAAGGAAGATGATGCTTTCAAACTCATTCTGAAAAAAA 415 410 tctttcagagaagtgttccaggacatgatgataagatacagtttgagtcttcactgtata 469 299 350 tttcctttaaggaaatgagtcctcctgagaatatcaatgatgaaggaaatgacatcatat 409 239 TTTCCTTTGAGGAAATGGATCCACCTGAAAATATTGATGATATACAAAGTGATCTCATAT 298 230 acgcaccccagaccgtatttatcatatatatgtataaagatagcctcactagaggtctag 289 119 GTGAACCCCAGACCAGACTGATAATATACATGTACAAAGACAGTGAAGTAAGAGGACTGG 178 170 tcattaaccagggaaatcaacctgtgtttgaggatatgcctgattctgattgtacagaca 229 62 TCGTTGACAAAAGA---CAGCCTGTGTTCGAGGATATGACTGATATTGATCAAAGTGCCA 118 2 ACTTTGGCCGACTTCACTGTACAACCGCAGTAATACGGAATATAAATGACCAAGTTCTCT 61 Local TCTTTCAGAAACGTGTTCCAGGAC---ACAACAAGATGGAGTTTGAATCTTCACTGTATG 355 Similarity Conservative 57.0%; 76.5%; 1, Pred. Score 268.4; DB Pred. No. 8.8e-67 Mismatches 102; Indels DB 21; Length 582; 6 Gaps

```
XX DE XX
                             A10526;
                                         A10526 standard; cDNA; 477 BP
```

Human interleukin-18 (IL-18) nucleotide sequence.

23-JUN-2000 (first entry)

Interleukin-18; production; IL-18; human; ss; medical injection product.

CN1243130-A

```
RESULT
V48226
                                                     맒
                                                                         Qy
                                                                                                     Дb
                                                                                                                             Qγ
                                                                                                                                                           Ъ
                                                                                                                                                                               Qy
                                                                                                                                                                                                             DЬ
                                                                                                                                                                                                                                     δÃ
                                                                                                                                                                                                                                                                  В
                                                                                                                                                                                                                                                                                      δÃ
                                                                                                                                                                                                                                                                                                                    Вþ
                                                                                                                                                                                                                                                                                                                                            Qy
                                                                                                                                                                                                                                                                                                                                                                        밁
                                                                                                                                                                                                                                                                                                                                                                                            Q
                                                                                                                                                                                                                                                                                                                                                                                                                          밁
                                                                                                                                                                                                                                                                                                                                                                                                                                                QY
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Query Match
Best Local S
Matches 346
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               stop codon preferred by coli bacillus to raise the expression rate method uses a high-amplification culture medium to increase the expression level and only requires a one-step purification process t
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Wu G,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Sequence
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        obtain a
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        sequence. The invention relates to a method for engineering bacterium for recombination of human II-18 and a method for the preparation of II-18. A primer containing a restriction endonuclease site can be used to accurately obtain the II-18 gene containing 474 nucleotides, and uses the
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              This sequence
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  interleukin-18
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Preparation method
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         P-PSDB;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                24-JUL-1998;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          24-JUL-1998;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   02~FEB-2000
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        (WUGG/)
                                                                               416
                                                     425
                                                                                                                                                                                                              245
                                                                                                                                   356
                                                                                                                                                             305
                                                                                                                                                                                       299
                                                                                                                                                                                                                                         239
                                                                                                                                                                                                                                                                   185
                                                                                                                                                                                                                                                                                                                       125
                                                                                                                                                                                                                                                                                                                                                                          65
                                                                                                                                                                                                                                                                                                                                                                                                                           5
                                                                                                                                                                                                                                                                                                                                                                                                                                                  2 ACTTTGGCCGACTTCACTGTACAACCGCAGTAATACGGAATATAAATGACCAAGTTCTCT
                                                                 AAGGACACTTTCTTGCTTGCCAAAAGGAAGATGATGCTTTCAAACTCATTCTGAAAAAAA
                                                                                                                                                                         TCTTTCAGAAACGTGTTCCAGGAC---ACAACAAGATGGAGTTTGAATCTTCACTGTATG
                                                                                                                                                                                                                                                                              CTGTGACCCTCTCTGTGAAGGATAGTAAAAAYGTCTACCCTCTCCTGTAAGAACAAGATCA
                                                                                                                                                                                                                                                                                                                   aggatgaattgggggatagatctataatgttcactgttcaaaacgaagactaa
                                                                                                                                                                                                              tttcctttaaggaaatgaatcctcctgataacatcaaggatacaaaaagtgacatcatat
                                                                                                                                                                                                                            TTTCCTTTGAGGAAATGGATCCACCTGAAAATATTGATGATATACAAAGTGATCTCATAT
                                                                                                                                                                                                                                                                 GTGAACCCCAGACCAGACTGATAATATACATGTACAAAGACAGTGAAGTAAGAGGACTGG
                                                                                                                                                                                                                                                                                                                                                                                     TCGTTGACAAAAGACA---GCCTGTGTTCGAGGATATGACTGATATTGATCAAAGTGCCA 118
                                                                                                                                                                                                                                                                                                                                                                                                                          actttggcaagcttgaatctaaatnatcagtcataagaaatttgaatgaccaagttctct
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       2000-340020/30
DB; Y85167.
                                                                                                      aaggatactttctagcttgtgaaaaagagagagacctttttaaactcattttgaaaaaag
                                                                                                                                                          tctttcagagaagtgtcccaggacatgataataagatgcaatttgaatcttcatcatacg
                                                                                                                                                                                                                                                                                                                                                                       tcattgaccaaggaaatcggcctctatttgaagatatgactgattctgactgtagagata
                                                                                                                                                                                                                                                                                                                                                                                                                                                                              346;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        æ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              477
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Page
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     medical injection-pure product.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              B₽;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       2;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            represents the human interleukin-18 (IL-18) nucleotide
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 and
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  98CN-0103307
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          98CN-0103307
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    17pp;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             169 A; 77
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 its
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             for
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          52.9%;
73.2%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              engineering bacteria for
product thereof -
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Ç
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Score 249.2; DB 2
Pred. No. 2.2e-61;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             89 G; 141 T; 1 other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Mismatches 120;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      DB 21;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             recombination
                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Length
                                                                                                                                                                                                                                                                                                                                                                                                                                                                              6
                                                                                                         424
                                                                                                                                                                                                              304
                                                                                                                                                                                                                                        298
                                                                                                                                                                                                                                                                  244
                                                                                                                                                                                                                                                                                                                    184
                                                                                                                                                                                                                                                                                                                                                                                                                          64
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ţ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                            2
```

¥₽

V48226

standard;

471

```
Qy
                                                   밁
                                                                                                                           Qy
                                                                          Qγ
                                                                                                    Ъ
                                                                                                                                                       В
                                                                                                                                                                            Qy
                                                                                                                                                                                                        Вþ
                                                                                                                                                                                                                             δÃ
                                                                                                                                                                                                                                                                                                                          Matches 344;
                                                                                                                                                                                                                                                                                                                                                           Interleukin-18 (IL-18) or a functional equivalent can be used for inhibition of osteoclast formation. IL-18 is used for treating or preventing osteoclast-related diseases e.g. hypercalcaemia, osteoclastoma Behcet's syndrome, osteosarcoma, arthropathy, chronic rheumatoid arthritis, deformity ostitis, primary hyperthyroidism, osteopenia and
                                                                                                                                                                                                                                                                                                                                                                                                                                                                          of e.g. hypercalcaemia, osteoclastoma, Behcet's syndrome, osteosarcoma, chronic rheumatoid arthritis, deformity ostitis.
                                                                                                                                                                                                                                                                                                                        Sequence 471 BP;
                                                                                                                                                                                                                                                                                                                                                osteoporosis.
                                                                                                                                                                                                                                                                                                                                                                                                                                      Disclosure; Page 19-20; 56pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Use of interleukin-18 to inhibit osteoclast formation - in treatment
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          WPI; 1998-448964/39
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Gillespie MT,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     25-FEB-1997;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             24-FEB-1998;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       02-SEP-1998
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           chronic
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        osteoclastoma
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     16-NOV-1998
 242
                                                                                                                             119
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            (наув
                                                                                                    122
                                                                                                                                                       62
                                                                                                                                                                                62
                                                                                                                                                                                                                                  N
             TTTCCTTTGAGGAAATGGATCCACCTGAAAATATTGATGATATACAAAGTGATCTCATAT
                                                              CTGTGACCCTCTCTGTGAAGGATAGTAAAAYGTCTACCCTCTCCTGTAAGAACAAGATCA
                                                                                                                  GTGAACCCCAGACCAGACTGATAATATACATGTACAAAGACAGTGAAGTAAGAGGACTGG
                                                                                                                                                                                                                     ACTTTGGCCGACTTCACTGTACAACCGCAGTAATACGGAATATAAATGACCAAGTTCTCT 61
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  sapiens
tttcctttaaggaaatgaatcctcctgataacatcaaggatacaaaaagtgacatcatat
                                                 tcattgaccaaggaaatcggcctctatttgaagatatgactgattctgactgtagagata
                                                                                                                                                                               TCGTTGACAAAAGACA---GCCTGTGTTCGAGGATATGACTGATATTGATCAAAAGTGCCA 118
                                                                                                                                                                                                      actttggcaagcttgaatctaaattatcagtcataagaaatttgaatgaccaagttctct 61
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        interleukin-18; IL-18; osteoclast; hypercalcaemia; osteopenia; lastoma Behcet's syndrome; osteosarcoma; arthropathy; osteoporo; rheumatoid arthritis; deformity ostitis; primary hyperthyroid:
                                                                                                                                                                                                                                                                       Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                              hyperthyroidism and osteoporosis
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               W77077.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            HAYASHIBARA
                                                                                                                                                                                                                                                           Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Horwood
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     97JP-0055468
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             98EP-0301352
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        /product=
/note= "No
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Location/Qualifiers
1..471
                                                                                                                                                                                                                                                                                                                       166 A;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               /*tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             18
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            SEIBUTSU KAGAKU
                                                                                                                                                                                                                                                                       52.6%;
73.3%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Ž,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         "No stop or start codon
                                                                                                                                                                                                                                                                                                                          77
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ื่อ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    "Interleukin
                                                                                                                                                                                                                                                                                                                        Ç
                                                                                                                                                                                                                                                           1;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Kurimoto
                                                                                                                                                                                                                                                                       Score 247.8; DB:
Pred. No. 5.4e-61;
                                                                                                                                                                                                                                                                                                                        88 G;
                                                                                                                                                                                                                                                           Mismatches 118;
                                                                                                                                                                                                                                                                                                                        140 T;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ĭ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Udagawa
                                                                                                                                                                                                                                                                                   DB 19;
                                                                                                                                                                                                                                                                                                                          0 other;
                                                                                                                                                                                                                                                           Indels
                                                                                                                                                                                                                                                                                  Length
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    steopenia; ds;
osteoporosis;
                                                                                                                                                                                                                                                           6,
                                                                                                                                                       121
                                                                          238
                                                                                                    181
                                                                                                                                                                                                                                                          Ŋ
```

```
Qy
                                                                                                                                                                                                                                                                                                                                                                                               14-NOV-1997;
29-NOV-1996;
21-JAN-1997;
The present sequence represents the mutant human interferon-gamma inducing factor cDNA IGIF/MUT12. The wild-type human interferon-gamma factor cDNA sequence is shown in V32754. The invention provides for mutant human and mouse interferon-gamma inducing factors in which one or more cysteine residues are replaced with different residues at or away from the consensus sequences shown in W48956-W48958. The mutant IGIFs are capable of stimulating immunocompetent cells for the production of interferon-gamma and are claimed to be less toxic, more active and
                                                                                                                                                                                                      Mutants of interferon-gamma inducing polypeptide - useful as antitumour, antiviral, antimicrobial or anti-immunopathic agents
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            mutation
                                                                                                                                                                   Claim 10; page 45; 59pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                        28-NOV-1997;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               EP845530-A2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                osteoporosis; thrombopenia; acquired immunodeficiency syndrome; ds.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Interferon-gamma inducing factor; interferon-gamma; killer cell; antitumour agent; tumour; antitumour agent; antiviral agent; antimicrobial agent; tumour; hepatitis; malaria; tuberculosis; renal carcinoma; rheumatism; A
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Mutant human interferon-gamma inducing factor cDNA IGIF/MUT12.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            25-SEP-1998 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      V32625 standard; cDNA; 471 BP
                                                                                                                                                                                                                                                                                                                      Kurimoto M, Okamoto I,
                                                                                                                                                                                                                                                                                                                                                        (HAYB ) HAYASHIBARA SEIBUTSU KAGAKU.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                422 aggatgaattgggggatagatctataatgttcactgttcaaaacgaaga 470
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     416 AGGATGAAAATGGGGATAAATCTGTAATGTTCACTCTCACTAACTTACA 464
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       362 aaggatactttctagcttgtgaaaaagagagagacctttttaaactcattttgaaaaaag 421
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              356 AAGGACACTTTCTTGCCTTGCCAAAAGGAAGATGATGCTTTCAAACTCATTCTGAAAAAAA 415
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      299 TCTTTCAGAAACGTGTTCCAGGAC---ACAACAAGATGGAGTTTGAATCTTCACTGTATG 355
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                302 tctttcagagaagtgtcccaggacatgataataagatgcaatttgaatcttcatcatacg 361
                                                                                                                                                                                                                                                                                 1998-288747/26.
                                                                                                                                                                                                                                                                                                                                                                                               97JP-0329715.
96JP-0333037.
97JP-0020906.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                        97EP-0309632.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       /note= "(
202..204
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 /note= "changed from TGT in wild-type to TCT in
    mutant"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             /product= "Mutant human interferon-gamma inducing
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           "CDS does not contain a stop codon"
                                                                                                                                                                                                                                                                                                                      Yamamoto K;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   rheumatism; AIDS;
```

```
δÃ
                                                                                                                                                                                                 Qy
                                                                                                                                                                                                                                                                                                  ρ
                                                                                                                                                                                                                                                                                                                                                       В
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       δÃ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             рь
                                                                                                                                                  В
                                                                                                                                                                                                                                                      밁
                                                                                                                                                                                                                                                                                                                                                                                                      Qγ
                                                                                                                                                                                                                                                                                                                                                                                                                                                           B
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            망
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           QY
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               QΥ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     stable than the corresponding wild type interferon-gamma inducing factor. The mutant IGIFs are also claimed to enhance killer cell cytotoxicity and/or induce killer cell formation, and may therefore be useful as antitumour agents, antitumour immunotherapeutics, antiviral agents and antimicrobial agents, The mutant IGIFs are also claimed to be useful for treating hepatitis, acquired immunodeficiency syndrome (AIDS), malaria, tuberculosis, solid malignant tumours (e.g. renal carcinoma), rheumatism, osteoporosis and thrombopenia caused by radiation- and chemo-therapy.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Sequence 471 BP; 166 A; 78 C; 87 G; 140 T; 0 other;
                                                                         356 AAGGACACTTTCTTGCCTAGAAAGGAAGATGATGCTTTCAAACTCATTCTGAAAAAAA 415
                                                                                                                                                        302
                                                                                                                                                                                                                                                                                                                                                                                    179 CTGTGACCCTCTCTGTGAAGGATAGTAAAAAYGTCTACCCTCTCCCTGTAAGAACAAGATCA 238
416 AGGATGAAAATGGGGATAAATCTGTAATGTTCACTCTCACTAACTTACA 464
                                                                                                                                                                                                                                                   242 tttcctttaaggaaatgaatcctcctgataacatcaaggatacaaaaagtgacatcatat 301
                                                                                                                                                                                                                                                                                                     239 TTTCCTTTGAGGAAATGGATCCACCTGAAAATATTGATGATATACAAAGTGATCTCATAT 298
                                                                                                                                                                                                                                                                                                                                                       182 ctgtaactatctctgtgaagtctgagaaaatttcaactctctcctgtgagaacaaaatta 241
                                                                                                                                                                                                                                                                                                                                                                                                                                                        62 TCGTTGACAAAAGACA---GCCTGTGTTCGAGGATATGACTGATATTGATCAAAGTGCCA 118
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            62 tcattgaccaaggaaatcggcctctatttgaagatatgactgattctgactgtagagata 121
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                2 actttggcaagcttgaatctaaattatcagtcataagaaatttgaatgaccaagttctct 61
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          2 ACTTTGGCCGACTTCACTGTACAACCGCAGTAATACGGAATATAAATGACCAAGTTCTCT 61
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Local
                                            aaggatactttctagcttgtgaaaaagagagagacctttttaaactcattttgaaaaaag
                                                                                                                                               tctttcagagaagtgtcccaggacatgataataagatgcaatttgaatcttcatcatacg 361
                                                                                                                                                                                                    TCTTTCAGAAACGTGTTCCAGGAC---ACAACAAGATGGAGTTTGAATCTTCACTGTATG 355
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             GTGAACCCCAGACCAGACTGATAATATACATGTACAAAGACAGTGAAGTAAGAGGACTGG 178
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     52.6%;
73.3%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Score 247.8; DB Pred. No. 5.4e-61
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Mismatches 118; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   DB 19; Length 471;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    6; Gaps
```

2

Search completed: November 25, 2000, 05:14:11 Job time: 2281 sec

422 aggatgaattgggggatagatctataatgttcactgttcaaaaacgaaga 470

	•		
τ			
•			
·			
•			
		•	